

in a decrease in one or more tumor markers, particularly a decrease in one or more serum tumor markers, in the mammal relative to baseline tumor marker levels.

Similarly, decreasing tumor marker concentrations or serum half lives after administration of the combination indicates a good prognosis, while tumor marker concentrations which decline slowly and do not reach the normal reference range predict residual tumor and poor prognosis. Further, during follow-up therapy, increases in tumor marker concentration predicts recurrent disease many months before clinical manifestation.

In addition to the above examples, Table No. 4, below, lists several references, hereby individually incorporated by reference herein, that describe tumor markers and their use in detecting and monitoring tumor growth and progression.

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Table No. 4. Tumor marker references.

European Group on Tumor Markers Publications Committee. Consensus Recommendations. Anticancer Research 19: 2785-2820 (1999)

Human Cytogenetic Cancer Markers. Sandra R. Wolman and Stewart Sell (eds.). Totowa, New Jersey: Humana Press. 1997

Cellular Markers of Cancer. Carleton Garrett and  
Stewart Sell (eds.). Totowa, New Jersey: Human Press.  
1995

Also included in the combination of the invention are  
the isomeric forms, prodrugs and tautomers of the  
5 described compounds and the pharmaceutically-acceptable  
salts thereof. Illustrative pharmaceutically acceptable  
salts are prepared from formic, acetic, propionic,  
succinic, glycolic, gluconic, lactic, malic, tartaric,  
citric, ascorbic, glucuronic, maleic, fumaric, pyruvic,  
10 aspartic, glutamic, benzoic, anthranilic, mesylic,  
stearic, salicylic, p-hydroxybenzoic, phenylacetic,  
mandelic,

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embonic (pamoic), methanesulfonic, ethanesulfonic, benzenesulfonic, pantothenic, toluenesulfonic, 2-hydroxyethanesulfonic, sulfanilic, cyclohexylaminosulfonic, algenic,  $\beta$ -hydroxybutyric, 5 galactaric and galacturonic acids. Suitable pharmaceutically-acceptable base addition salts of compounds of the present invention include metallic ion salts and organic ion salts. More preferred metallic ion salts include, but are not limited to appropriate alkali 10 metal (group Ia) salts, alkaline earth metal (group IIa) salts and other physiological acceptable metal ions. Such salts can be made from the ions of aluminum, calcium, lithium, magnesium, potassium, sodium and zinc. Preferred organic salts can be made from tertiary amines and 15 quaternary ammonium salts, including in part, trimethylamine, diethylamine, N,N'-dibenzylethylenediamine, chloroprocaine, choline, diethanolamine, ethylenediamine, meglumine (N-methylglucamine) and procaine. All of the above salts can 20 be prepared by those skilled in the art by conventional means from the corresponding compound of the present invention.

Administration Regimen

25 Any effective treatment regimen can be utilized and readily determined and repeated as necessary to effect treatment. In clinical practice, the compositions containing an COX-2 inhibitor alone or in combination with other therapeutic agents are administered in 30 specific cycles until a response is obtained.